

**CENTER FOR DRUG EVALUATION AND
RESEARCH**

APPLICATION NUMBER: 20-986

ENVIRONMENTAL ASSESSMENT AND/OR FONSI

3.5 Environmental Assessment

A categorical exclusion from submitting an environmental assessment for the drug product, Insulin aspart, is requested based on the regulations in 21 CFR, part 25, Section 25.31. Section 25.31b provides for categorical exclusion if the concentration of the drug substance at the point of entry into the aquatic environment is below 1 part per billion (1 ppb).

Using the method of calculation for the Expected Introduction Concentration (EIC) of the drug as given in the "Guidance for Industry -- Environmental Assessment of Human Drug and Biologics Applications" (CDER/CBER, FDA July 1998), it is estimated that the EIC for Insulin aspart is _____ ppb and to the best of our knowledge, no extraordinary circumstances exist (21 CFR, 25.15(d)). This amount of drug substance does not exceed the EIC of 1 ppb and therefore the request is in compliance with the regulations for categorical exclusion. The calculation is described below.

EIC:

Assumptions for calculation:

1. Amount of drug per year:
2. Even daily distribution throughout the United States
3. No metabolism or depletion mechanism

Formula: $EIC = (A)(B)(C)(D)$ where

A = kg/year production _____ year)

B = _____ 'day entering Publicly Owned Wastewater Treatment Plants)

C = Year/365 days

D = _____

EIC = _____ ppb

**APPEARS THIS WAY
ON ORIGINAL**